

# Assessing Uncertainty in High-dimensional Regression Models

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# Papers:

- ▶ Statistical significance in high-dimensional linear models, Bühlmann (2012).
- ▶ On asymptotically optimal confidence regions and tests for high-dimensional models, van de Geer, Bühlmann, and Ritov (2013).

# Associations: Marginal v.s. Conditional

Genes:  $\{A^*\}$   $\{B_1^*, B_2, B_3\}$   $\{C_1, C_2, C_3\}$

(An example from Grazier G'Sell et al. 2013. )

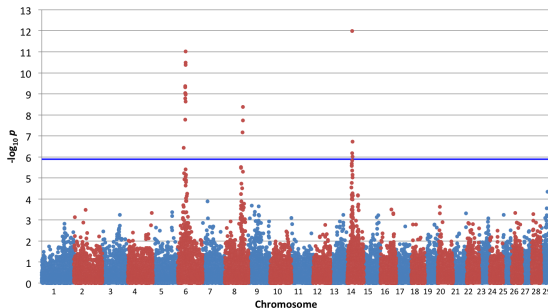


Figure: A Manhattan plot from Nishimura et al. (2012).

# Regression models on high-dimensional data

$$Y = X\beta^* + \epsilon, \quad \epsilon \sim N_n(0, I_n). \quad (1)$$

OLS/ML on this simple model is equivalent to:

$$\underset{\beta \in \mathbb{R}^p}{\text{minimize}} \quad \|Y - X\beta\|_2^2. \quad (2)$$

Linear algebra gives  $\hat{\beta} = (X^T X)^{-1} X^T Y$ .

However,  $X^T X$  is non-invertible when  $p > n$ . Consider the ridge regression:

$$\underset{\beta \in \mathbb{R}^p}{\text{minimize}} \quad \|Y - X\beta\|_2^2 + \lambda_n \sum_{i=1}^p \beta_i^2. \quad (3)$$

# Penalized regression

There are many choices of penalties:

- ▶  $\ell_2$  norm, i.e.  $\|\beta\|_2^2 = \sum_{i=1}^p \beta_i^2$ , ridge regressions.
- ▶  $\ell_1$  norm, i.e.  $\|\beta\|_1 = \sum_{i=1}^p |\beta_i|$ , lasso regressions.
- ▶ and more.

Advantages of using certain penalties:

1. Each penalty imposes a **low-dimensional** structure.
2. **Convex** loss functions.

# Assessing uncertainty

- ▶ Lasso: “Sparse estimators such as Lasso do not have a tractable limiting distribution. The limiting distributions of Lasso estimators **depend on unknown parameters** in low-dimensional setting. Bootstrap and subsampling techniques are plagued by **non-continuity** of limiting distributions.” (van de Geer et al., 2013).
- ▶ Ridge regression:  $\text{Bias}(\hat{\beta}) = -\lambda_n(X^T X + \lambda_n I)^{-1} \beta$ ; Bootstrap in low-dimensional setting, see Crivelli et al. (1995).

- ▶ Variable selections: Correct recover of support requires the “[beta-min](#)” assumption (Meinshausen and Bühlmann 2006, Wainwright 2009, Negahban et al. 2010).
- ▶ Bootstrap and subsampling method. In [low-dimensional setting](#), bootstraps for lasso are proposed by Chatterjee and Lahiri (2011), Sartori (2011). For Ridge regression, see Crivelli et al. (1995).
- ▶ A significance test for the lasso (Lockhart, Taylor, Tibshirani, and Tibshirani, 2013).

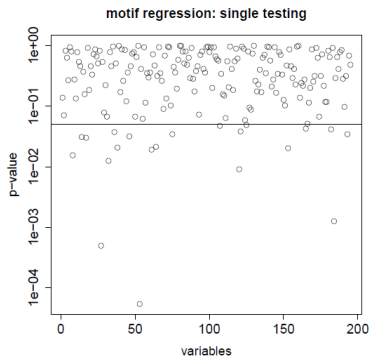
## Another line of research...

- ▶ Zhang and Zhang (2011), and Bühlmann (2012) provided hypothesis testing procedures on high-dimensional linear models.
- ▶ Javanmard and Montanari (2013) provided a minimax test for linear models.
- ▶ van de Geer et al. (2013) claimed to reach the **semiparametric efficiency bound**, and the method be generalized to  $\ell_1$ -penalized **GLMs** and ridge regressions.

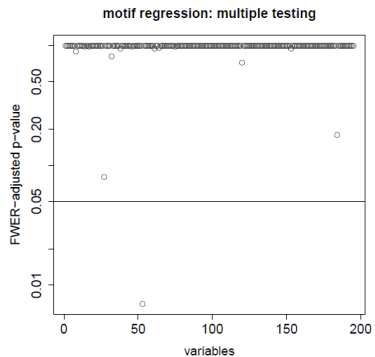
# Summary

- ▶ Marginal associations v.s. conditional associations.
- ▶ Reasons for using penalized regressions on high-dimensional data.
- ▶ Current attempts to make statistical inference on high-dimensional regressions.

# A glimpse ahead



(a)



(b)

Figure: A Motif regression result from Bühlmann (2012).

# The method in van de Geer et al. (2013)

$$\hat{\beta} = \underset{\beta \in \mathbb{R}^p}{\operatorname{argmin}} (\|Y - X\beta\|_2^2 / (2n) + \lambda \|\beta\|_1). \quad (4)$$

The Karush-Kuhn-Tucker conditions are

$$\begin{aligned} -X^T(Y - X\hat{\beta}) + \lambda\hat{\tau} &= 0, \\ \|\hat{\tau}\|_\infty &\leq 1, \text{ and } \hat{\tau}_j = \operatorname{sign}(\hat{\beta}_j) \text{ if } \hat{\beta}_j \neq 0. \end{aligned} \quad (5)$$

Using the first equation, we have

$$n^{-1}X^T X(\hat{\beta} - \beta^*) + \lambda\hat{\tau} = X^T \epsilon / n. \quad (6)$$

Now assume we have a  $\hat{\Theta}$  that is a “relaxed form” of an inverse of  $n^{-1}X^T X$ .

$$\hat{\beta} - \beta^* + \hat{\Theta}\lambda\hat{\tau} = \hat{\Theta}X^T \epsilon / n - \Delta, \quad (7)$$

where  $\Delta = (\hat{\Theta}\hat{\Sigma} - I)(\hat{\beta} - \beta)$ .

# The method in van de Geer et al. (2013), continued

We will show that  $\Delta$ , in fact  $\sqrt{n}\Delta$ , is asymptotically negligible under certain assumptions on sparsity. Then let

$$\hat{b} = \hat{\beta} + \hat{\Theta}X^T(Y - X\hat{\beta})/n. \quad (8)$$

Theorem 2.2 in van de Geer et al. (2013) claims that:

$$\sqrt{n}(\hat{b} - \beta^*) = W + o_P(1), \quad W|X \sim N_p(0, \sigma_\epsilon^2 \hat{\Theta} \hat{\Sigma} \hat{\Theta}^T). \quad (9)$$

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